

Amendment

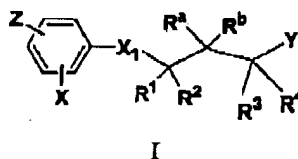
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USSN 09/848,697
QA211NPAmendments to the Claims:

This listing of claims will replace all prior versions and listings of claims in the application:

Listing of Claims:

Claim 1. (Currently Amended) A compound of the formula



wherein X_1 is O, $S(O)_n$, $-N^{R^5}-$, $CO-N^{R^6}-$, or $-CH_2-$, with the proviso that when X_1 is $-CH_2-$, R^1 and R^2 are only halogen;

n is 0, 1 or 2;

R^a and R^b when taken together form an oxo ($=O$) group, or R^a and R^b are each independently hydrogen, OH, $OCOR^9$, NH_2 , N_3 , $NHCOOR^9$, $NHCOCOR^9$, $NHSO_2R^9$ or F;

X is H, CF_3 , OCF_3 , halogen, C_1-C_7 alkyl, C_2-C_7 alkenyl, C_2-C_7 alkynyl or C_3-C_7 cycloalkyl, said alkyl, alkenyl, alkynyl or cycloalkyl group being optionally substituted by $COOR^8$, CN, $C(O)NR^6R^7$, PO_3R^8 , SO_3R^8 , heterocyclic, OR^8 , SH, $S(O)_nR^9$, NR^6R^7 , $NH(CO)NR^6R^7$, $NH(CO)OR^9$, or aryl or heteroaryl, said aryl or heteroaryl being optionally substituted by one or two groups independently selected from NR^6R^7 , OR^8 , $COOR^8$, SO_3R^8 , $OCOR^9$, PO_3R^8 , and $C(O)NR^6R^7$ and heterocyclic;

R^1 and R^2 are each independently H, halogen, OR^9 , C_1-C_7 alkyl, C_2-C_7 alkynyl,

O.K. to enter Sp. 10/19/04

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C₂-C₇ alkenyl or C₃-C₇ cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by COOR⁸, CN, C(O)NR⁶R⁷, PO₃R⁸, SO₃R⁸, heterocyclic, OR⁸, SH, S(O)_nR⁹, NR⁶R⁷, NH(CO)NR⁶R⁷, NH(CO)OR⁹, OC(O)OR⁹, or aryl or heteroaryl, said aryl and heteroaryl being optionally substituted with one or two groups independently selected from NR⁶R⁷, OR⁸, COOR⁸, SO₃R⁸, OCOR⁹, PO₃R⁸, and C(O)NR⁶R⁷ and heterocyclic;

R³, R⁴ and Y are each independently H, halogen, OR¹⁰, S(O)_nR¹⁰, C₁-C₇ alkyl, C₂-C₇ alkenyl, C₂-C₇ alkynyl or C₃-C₇ cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by COOR⁸, CN, C(O)NR⁶R⁷, PO₃R⁸, SO₃R⁸, heterocyclic, OR⁸, SH, S(O)_nR⁹, NR⁶R⁷, NH(CO)NR⁶R⁷, NH(CO)OR⁹, OC(O)OR⁹, or aryl or heteroaryl, said aryl and heteroaryl being optionally substituted by one or two groups independently selected from NR⁶R⁷, OR⁸, COOR⁸, SO₃R⁸, OCOR⁸, PO₃R⁸, and C(O)NR⁶R⁷ and heterocyclic, with the proviso that not all of R³, R⁴ and Y may be the same halogen;

R⁵, R⁶ and R⁷ are each independently H, C₁-C₇ alkyl, C₂-C₇ alkenyl, C₂-C₇ alkynyl or C₃-C₇ cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by COOR⁸, CN, OR⁸, NR⁸R⁹, SO₃R⁸, PO₃R⁸, halogen, or aryl or heteroaryl, said aryl or heteroaryl being optionally substituted by one or two groups independently selected from COOR⁸, SO₃R⁸, and PO₃R⁸ and heterocyclic;

R⁸ is H, C₁-C₇ saturated straight chain alkyl or cycloalkyl;

R⁹ is C₁-C₇ saturated straight chain alkyl or cycloalkyl;

R¹⁰ is C₁-C₇ alkyl, C₂-C₇ alkenyl, C₂-C₇ alkynyl, aryl or C₃-C₇ cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being optionally substituted by

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COOR⁸, CN, C(O)NR⁶R⁷, PO₃R⁸, SO₃R⁸, heterocyclic, OR⁸, SH, S(O)_nR⁹,
NR⁶R⁷, NH(CO)NR⁶R⁷, NH(CO)OR⁹, or aryl or heteroaryl, said aryl or heteroaryl
being optionally substituted by one or two groups independently selected from
NR⁶R⁷, OR⁸, COOR⁸, SO₃R⁸, OCOR⁸, PO₃R⁸, and C(O)NR⁶R⁷ and heterocyclic;

Z is OR¹¹, S(O)_nR¹¹, NR¹¹R¹² or CHR¹¹R¹²;

R¹¹ is C₁-C₇ alkyl, C₂-C₇ alkenyl, C₂-C₇ alkynyl or C₃-C₇ cycloalkyl, said alkyl,
alkenyl, alkynyl or cycloalkyl group being substituted by NR¹³R¹⁴, S(O)_nR¹³, or
OR¹³;

R¹² is hydrogen, C₁-C₇ alkyl, C₂-C₇ alkenyl, C₂-C₇ alkynyl or C₃-C₇ cycloalkyl,
said alkyl, alkenyl, alkynyl or cycloalkyl group being optionally substituted by
NR¹³R¹⁴, S(O)_nR¹³, or OR¹³;

R¹³ is ~~SiR¹⁵R¹⁶R¹⁷~~, C₁-C₇ alkyl, C₂-C₇ alkenyl, C₂-C₇ alkynyl, aryl or C₃-C₇
cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being substituted by
one to three groups independently selected from COOR⁸, OR⁸, ~~SiR¹⁵R¹⁶R¹⁷~~,
OR¹⁵, aryl, and biaryl and heteroaryl, said aryl[[.]] and biaryl and heteroaryl being
optionally substituted with one to three groups independently selected from halogen,
CF₃, OR⁸, COOR⁸, NO₂, and CN;

R¹⁴ is H, ~~SiR¹⁵R¹⁶R¹⁷~~, C₁-C₇ alkyl, C₂-C₇ alkenyl, C₂-C₇ alkynyl, aryl or C₃-
C₇ cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being optionally
substituted by one to three groups independently selected from COOR⁸, OR⁸, ~~SiR¹⁵R¹⁶R¹⁷~~,
~~R¹⁵R¹⁶R¹⁷~~, OR¹⁵, aryl, and biaryl and heteroaryl, said aryl[[.]] and biaryl and
heteroaryl being optionally substituted with one to three groups independently
selected from halogen, CF₃, OR⁸, COOR⁸, NO₂, and CN; and ~~or~~

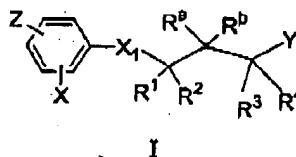
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~~R¹³ and R¹⁴ when taken together with the nitrogen atom to which they are attached may form a 5-7 membered heterocyclic ring with one or more heteroatoms selected from O, N and S; said ring being optionally substituted by OR⁸, COOR⁸, or C(O)NR⁶R⁷, and~~

R¹⁵, R¹⁶, R¹⁷ are each independently is C₁-C₇ alkyl, aryl, benzyl, benzhydryl, biaryl, heteroaryl, or (C₁-C₆) alkyl-aryl or (C₁-C₆) alkyl-heteroaryl, said aryl, benzyl, benzhydryl, and biaryl being optionally substituted by halogen, CF₃, OR⁸, COOR⁸, NO₂, CN, or C₁-C₇ alkyl.

Claim 2. (Currently Amended) A compound of the formula



or a pharmaceutically acceptable salt thereof wherein

X₁ is O, S(O)_n, $\text{--}\overset{\text{R}^5}{\text{N}}\text{--}$, $\text{CO--}\overset{\text{R}^6}{\text{N}}\text{--}$ or $\text{--CH}_2\text{--}$, with the proviso that when X₁ is $\text{--CH}_2\text{--}$, R¹ and R² are only halogen;

n is 0, 1 or 2;

R^a and R^b when taken together form an oxo (=O) group, or R^a and R^b are each independently hydrogen, OH, OCOR⁹, NH₂, N₃, NHCOOR⁹, NHCOCOR⁹, NHSO₂R⁹ or F;

X is H, CF₃, OCF₃, halogen, C₁-C₇ alkyl, C₂-C₇ alkenyl, C₂-C₇ alkynyl or C₃-C₇ cycloalkyl, said alkyl, alkenyl, alkynyl or cycloalkyl group being optionally substituted by COOR⁸, CN, C(O)NR⁶R⁷, PO₃R⁸, SO₃R⁸, heterocyclic, OR⁸, SH,

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$S(O)_nR^9$, NR^6R^7 , $NH(CO)NR^6R^7$, $NH(CO)OR^9$, or aryl or heteroaryl, said aryl or heteroaryl being optionally substituted by one or two groups independently selected from NR^6R^7 , OR^8 , $COOR^8$, SO_3R^8 , $OCOR^9$, PO_3R^8 , and $C(O)NR^6R^7$ and heterocyclo;

R^1 and R^2 are each independently H, halogen, OR^9 , C_1-C_7 alkyl, C_2-C_7 alkynyl, C_2-C_7 alkenyl or C_3-C_7 cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by $COOR^8$, CN, $C(O)NR^6R^7$, PO_3R^8 , SO_3R^8 , heterocyclo, OR^8 , SH, $S(O)_nR^9$, NR^6R^7 , $NH(CO)NR^6R^7$, $NH(CO)OR^9$, $OC(O)OR^9$, or aryl or heteroaryl, said aryl and heteroaryl being optionally substituted with one or two groups independently selected from NR^6R^7 , OR^8 , $COOR^8$, SO_3R^8 , $OCOR^9$, PO_3R^8 , and $C(O)NR^6R^7$ and heterocyclo;

R^3 , R^4 and Y are each independently H, OR^{10} , $S(O)_nR^{10}$, C_1-C_7 alkyl, C_2-C_7 alkynyl, C_2-C_7 alkenyl or C_3-C_7 cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by $COOR^8$, CN, $C(O)NR^6R^7$, PO_3R^8 , SO_3R^8 , heterocyclo, OR^8 , SH, $S(O)_nR^9$, NR^6R^7 , $NH(CO)NR^6R^7$, $NH(CO)OR^9$, $OC(O)OR^9$, or aryl or heteroaryl, said aryl and heteroaryl being optionally substituted by one or two groups independently selected from NR^6R^7 , OR^8 , $COOR^8$, SO_3R^8 , $OCOR^8$, PO_3R^8 , and $C(O)NR^6R^7$ and heterocyclo;

R^5 , R^6 and R^7 are each independently H, C_1-C_7 alkyl, C_2-C_7 alkenyl, C_2-C_7 alkynyl or C_3-C_7 cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by $COOR^8$, CN, OR^8 , NR^8R^9 , SO_3R^8 , PO_3R^8 , halogen, or aryl or heteroaryl, said aryl and heteroaryl being optionally substituted by one or two groups independently selected from $COOR^8$, SO_3R^8 , and PO_3R^8 and heterocyclo;

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R^8 is H, C₁-C₇ saturated straight chain alkyl or cycloalkyl, CF₃ or CH₂CF₃;

R^9 is C₁-C₇ saturated straight chain alkyl or cycloalkyl;

R^{10} is C₁-C₇ alkyl, C₂-C₇ alkenyl, C₂-C₇ alkynyl, aryl or C₃-C₇ cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being optionally substituted by COOR⁸, CN, C(O)NR⁶R⁷, PO₃R⁸, SO₃R⁸, heterocyclic, OR⁸, SH, S(O)_nR⁹, NR⁶R⁷, NH(CO)NR⁶R⁷, NH(CO)OR⁹, or aryl or heteroaryl, said aryl or heteroaryl being optionally substituted by one or two groups independently selected from NR⁶R⁷, OR⁸, COOR⁸, SO₃R⁸, OCOR⁸, PO₃R⁸, and C(O)NR⁶R⁷ and heterocyclic;

Z is OR¹¹, S(O)_nR¹¹, NR¹¹R¹² or CHR¹¹R¹²;

R^{11} is C₁-C₇ alkyl, C₂-C₇ alkenyl, C₂-C₇ alkynyl or C₃-C₇ cycloalkyl, said alkyl, alkenyl, alkynyl or cycloalkyl group being substituted by NR¹³R¹⁴, S(O)_nR¹³, or OR¹³;

R^{12} is hydrogen, C₁-C₇ alkyl, C₂-C₇ alkenyl, C₂-C₇ alkynyl or C₃-C₇ cycloalkyl, said alkyl, alkenyl, alkynyl or cycloalkyl group being optionally substituted by NR¹³R¹⁴, S(O)_nR¹³ or OR¹³;

R^{13} is SiR¹⁵R¹⁶R¹⁷, C₁-C₇ alkyl, C₂-C₇ alkenyl, C₂-C₇ alkynyl, aryl or C₃-C₇ cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being substituted by one to three groups independently selected from COOR⁸, OR⁸, SiR¹⁵R¹⁶R¹⁷, OR¹⁵, aryl, and biaryl and heteroaryl, said aryl[[.]] and biaryl and heteroaryl being optionally substituted with one to three groups independently selected from halogen, CF₃, OR⁸, COOR⁸, NO₂, and CN;

R^{14} is H, SiR¹⁵R¹⁶R¹⁷, C₁-C₇ alkyl, C₂-C₇ alkenyl, C₂-C₇ alkynyl, aryl or C₃-

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C₇ cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being optionally substituted by one to three groups independently selected from COOR⁸, OR⁸, SiR¹⁵R¹⁶R¹⁷, OR¹⁵, aryl, and biaryl and heteroaryl, said aryl[[,]] and biaryl and heteroaryl being optionally substituted with one to three groups independently selected from halogen, CF₃, OR⁸, COOR⁸, NO₂, and CN; and or

~~R¹³ and R¹⁴ when taken together with the nitrogen atom to which they are attached may form a 5-7 membered heterocyclic ring with one or more heteroatoms selected from O, N and S; said ring being optionally substituted by OR⁸, COOR⁸, or C(O)NR⁵R⁶; and~~

R¹⁵, ~~R¹⁶, R¹⁷~~ are each independently is C₁-C₇ alkyl, aryl, benzyl, benzhydryl, biaryl, heteroaryl, or (C₁-C₆) alkyl-aryl or (C₁-C₆) alkyl-heteroaryl, said aryl, benzyl, benzhydryl, and biaryl being optionally substituted by halogen, CF₃, OR⁸, COOR⁸, NO₂, CN, or C₁-C₇ alkyl.

Claim 3. (Currently Amended) A compound of claim 2 wherein X₁ is O or S(O)_n and Y is OR¹⁰ in which R¹⁰ is C₁-C₇ alkyl, C₂-C₇ alkenyl, C₂-C₇ alkynyl, aryl or C₃-C₇ cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being optionally substituted by COOR⁸, CN, C(O)NR⁶R⁷, PO₃R⁸, SO₃R⁸, heterocyclic, OR⁸, SH, S(O)_nR⁹, NR⁶R⁷, NH(CO)NR⁶R⁷, NH(CO)OR⁹, or aryl or heteroaryl, said aryl or heteroaryl being optionally substituted by one or two groups independently selected from NR⁶R⁷, OR⁸, COOR⁸, SO₃R⁸, OCOR⁹, PO₃R⁸, and C(O)NR⁶R⁷ or heterocyclic, said R⁶, R⁷, R⁸ and R⁹ substituents being defined as in claim 2.

Claim 4. (Original) A compound of claim 3 in which R^a and R^b taken together represent an oxo (=O) group, or R^a and R^b are each independently hydrogen or OH.

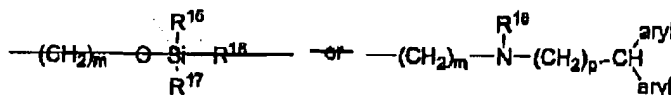
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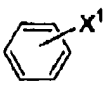
Claims 5-6. (Canceled).

Claim 7. (Currently Amended) A compound of claim 3 in which

Z is



in which m and p each independently represent an integer of one to six, R^{15} , R^{16} , R^{17} are each independently C_1 - C_7 alkyl or phenyl, R^{18} is C_1 - C_7 alkyl and aryl

represents  in which X^1 is halogen.

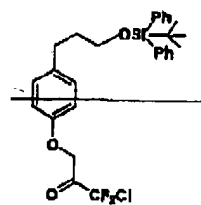
Claim 8. (Canceled).

Claim 9. (Original) A pharmaceutical composition for the inhibition of cytosolic phospholipase A_2 comprising a therapeutically effective amount of a compound of claim 1 and a pharmaceutically acceptable carrier.

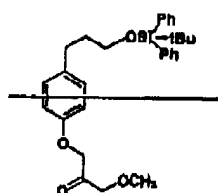
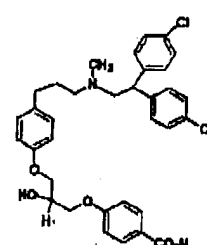
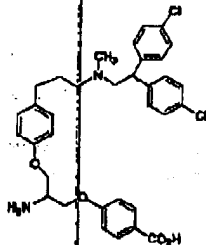
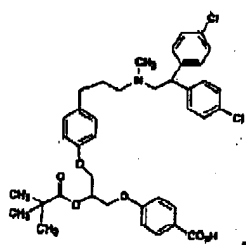
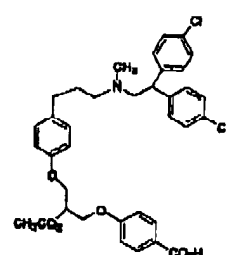
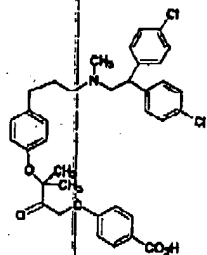
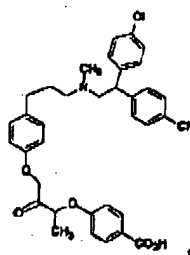
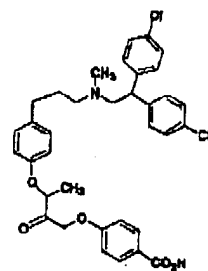
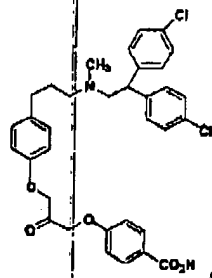
Claim 10. (Withdrawn) A method of inhibiting cytosolic phospholipase A_2 in a mammal in need thereof, comprising administering to said mammal a therapeutically effective amount of a compound of claim 1.

Claim 11. (Currently Amended) A compound selected from

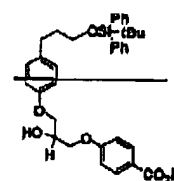
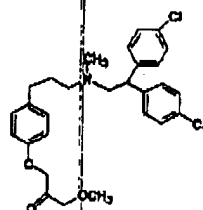
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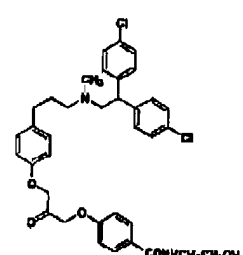
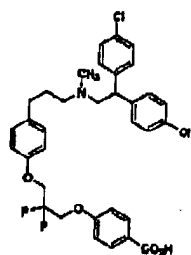
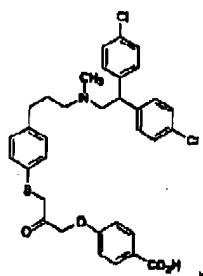
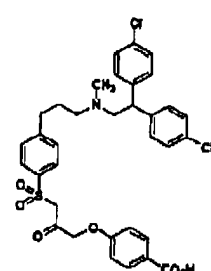
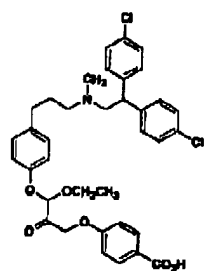
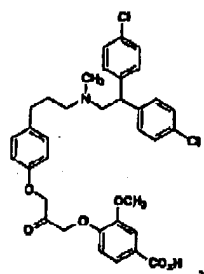
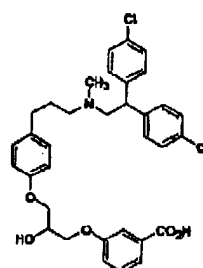
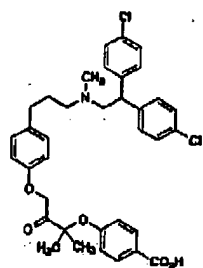
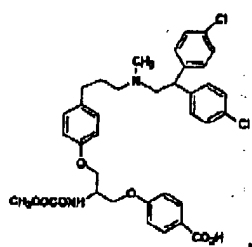
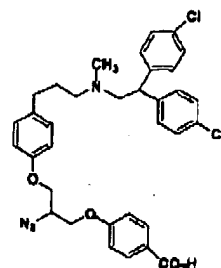
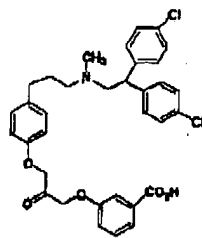
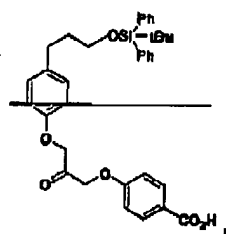


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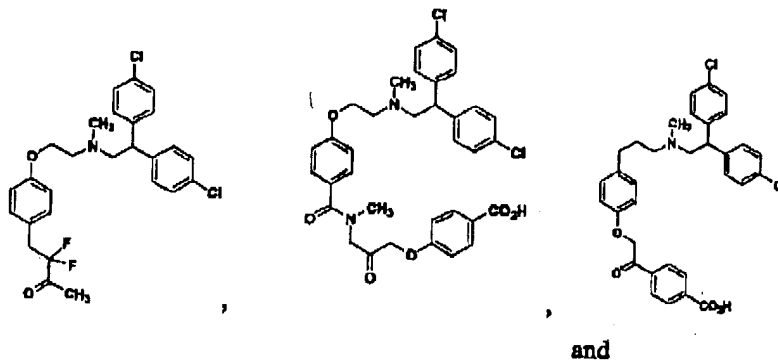


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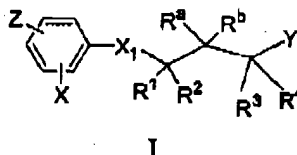
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or a pharmaceutically acceptable salt thereof.

Claim 12. (Currently Amended) A compound of the formula



or a pharmaceutically acceptable salt thereof wherein

X_1 is O, $S(O)_n$, $CO-N^{R^5}-$, or $-CH_2-$, with the proviso that when X_1 is $-CH_2-$, R^1 and R^2 are only halogen;

n is 0, 1 or 2;

R^a and R^b when taken together form an oxo ($=O$) group, or R^a and R^b are each independently hydrogen, OH, $OCOR^3$, NH_2 , N_3 , $NHCOCOR^3$, or F;

X is H;

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R^1 and R^2 are each independently H, halogen, OR^9 , or C_1-C_7 alkyl;

R^3 , R^4 and Y are each independently H, halogen, OR^{10} , or C_1-C_7 alkyl, said alkyl being optionally substituted by aryl, said aryl being optionally substituted by one or two $COOR^8$ groups, with the proviso that not all of R^3 , R^4 and Y may be the same halogen;

R^5 , R^6 , and R^7 are each independently hydrogen or C_1-C_7 alkyl, said alkyl being optionally substituted by OR^8 ;

R^8 is H or C_1-C_7 saturated straight chain alkyl;

R^9 is C_1-C_7 saturated straight chain alkyl;

R^{10} is C_1-C_7 alkyl or aryl, said alkyl or aryl group being optionally substituted by $COOR^8$, $C(O)NR^6R^7$, heterocyclic, or OR^8 ;

Z is OR^{11} or $CHR^{11}R^{12}$;

R^{11} is C_1-C_7 alkyl substituted by $NR^{13}R^{14}$, $S(O)_nR^{13}$, or OR^{13} ;

R^{12} is hydrogen;

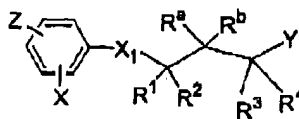
R^{13} is ~~$SiR^{15}R^{16}R^{17}$~~ or C_1-C_7 alkyl, said alkyl substituted by one to three groups independently selected from OR^{15} and aryl, said aryl substituted with one halogen;

R^{14} is C_1-C_7 alkyl; and

~~R^{15} , R^{16} , and R^{17}~~ are each independently is C_1-C_7 alkyl, aryl, or benzhydryl, said aryl and benzhydryl being optionally substituted by halogen.

Claim 13. (Currently Amended) A compound of the formula

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or a pharmaceutically acceptable salt thereof wherein

X_1 is O, $S(O)_n$, or $-CH_2-$, with the proviso that when X_1 is $-CH_2-$, R^1 and R^2 are only halogen;

n is 0, 1 or 2;

R^a and R^b are each independently hydrogen, OH, $OCOR^9$, NH_2 , N_3 , $NHCOOR^9$, $NHCOCOR^9$, or F;

X is H, CF_3 , OCF_3 , halogen, C_1-C_7 alkyl, C_2-C_7 alkenyl, C_2-C_7 alkynyl or C_3-C_7 cycloalkyl, said alkyl, alkenyl, alkynyl or cycloalkyl group being optionally substituted by $COOR^8$, CN, $C(O)NR^6R^7$, PO_3R^8 , SO_3R^8 , heterocyclic, OR^8 , SH, $S(O)_nR^9$, NR^6R^7 , $NH(CO)NR^6R^7$, $NH(CO)OR^9$, or aryl or heteroaryl; said aryl or heteroaryl being optionally substituted by one or two groups independently selected from NR^6R^7 , OR^8 , $COOR^8$, SO_3R^8 , $OCOR^9$, PO_3R^8 , and $C(O)NR^6R^7$ and heterocyclic;

R^1 and R^2 are each independently H, halogen, OR^9 , C_1-C_7 alkyl, C_2-C_7 alkynyl, C_2-C_7 alkenyl or C_3-C_7 cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by $COOR^8$, CN, $C(O)NR^6R^7$, PO_3R^8 , SO_3R^8 , heterocyclic, OR^8 , SH, $S(O)_nR^9$, NR^6R^7 , $NH(CO)NR^6R^7$, $NH(CO)OR^9$, $OC(O)OR^9$, or aryl or heteroaryl, said aryl and heteroaryl being optionally substituted with one or two groups independently selected from NR^6R^7 , OR^8 , $COOR^8$, SO_3R^8 , $OCOR^9$, PO_3R^8 , and $C(O)NR^6R^7$ and heterocyclic;

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R^3 and R^4 are each independently H, halogen, OR^{10} , $S(O)_nR^{10}$, C_1 - C_7 alkyl, C_2 - C_7 alkenyl, C_2 - C_7 alkynyl or C_3 - C_7 cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by $COOR^8$, CN, $C(O)NR^6R^7$, PO_3R^8 , SO_3R^8 , ~~heterocyclic~~, OR^8 , SH, $S(O)_nR^9$, NR^6R^7 , $NH(CO)NR^6R^7$, $NH(CO)OR^9$, $OC(O)OR^9$, ~~or aryl or heteroaryl~~, said aryl and heteroaryl being optionally substituted by one or two groups independently selected from NR^6R^7 , OR^8 , $COOR^8$, SO_3R^8 , $OCOR^8$, PO_3R^8 , and $C(O)NR^6R^7$ and heterocyclic, ~~with the proviso that not all of R^3 , R^4 and Y may be the same halogen.~~

Y is OR^{10} or $S(O)_nR^{10}$;

R^5 , R^6 and R^7 are each independently H, C_1 - C_7 alkyl, C_2 - C_7 alkenyl, C_2 - C_7 alkynyl or C_3 - C_7 cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by $COOR^8$, CN, OR^8 , NR^8R^9 , SO_3R^8 , PO_3R^8 , halogen, ~~or aryl or heteroaryl~~, said aryl ~~or heteroaryl~~ being optionally substituted by one or two groups independently selected from $COOR^8$, SO_3R^8 , and PO_3R^8 and heterocyclic;

R^8 is H, C_1 - C_7 saturated straight chain alkyl or cycloalkyl;

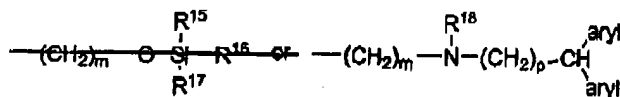
R^9 is C_1 - C_7 saturated straight chain alkyl or cycloalkyl;

R^{10} is C_1 - C_7 alkyl, C_2 - C_7 alkenyl, C_2 - C_7 alkynyl, aryl or C_3 - C_7 cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being optionally substituted by $COOR^8$, CN, $C(O)NR^6R^7$, PO_3R^8 , SO_3R^8 , ~~heterocyclic~~, OR^8 , SH, $S(O)_nR^9$, NR^6R^7 , $NH(CO)NR^6R^7$, $NH(CO)OR^9$, ~~or aryl or heteroaryl~~, said aryl ~~or heteroaryl~~ being optionally substituted by one or two groups independently selected from NR^6R^7 , OR^8 , $COOR^8$, SO_3R^8 , $OCOR^8$, PO_3R^8 , and $C(O)NR^6R^7$ or heterocyclic;
and

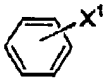
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Z is



in which m and p each independently represent an integer of one to six, R^{15} , R^{16} , R^{17} are each independently C_1 - C_7 alkyl or phenyl, R^{18} is C_1 - C_7 alkyl and aryl

represents  in which X^1 is halogen.

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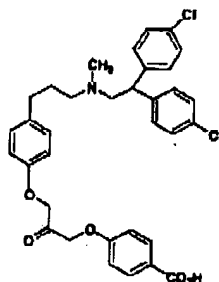
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QA211NPREMARKS/ARGUMENTS

Claims 1-4, 7, and 9-13 are pending in this application.

In the Office Action dated July 8, 2004, the Examiner rejected Claims 1-4, 9, 12, and 13 under 35 U.S.C. §102(b) as being unpatentable over JP 5-222006. Claim 7 was objected to as being dependent upon a rejected base claim.

Reconsideration and allowance of this application are respectfully requested in view of the above amendments and the remarks that follow.

Pursuant to the Examiner's request for an election of a single disclosed species on July 29, 2003, Applicants elected 3-[4-[3-[N-[2-Bis-(4-chlorophenyl)ethyl]-N-methylamino]propyl]phenoxy]-1-(4-carboxyphenoxy)-2-propanone which is Example 2 on page 51 of the specification.



3-[4-[3-[N-[2-Bis-(4-chlorophenyl)ethyl]-N-methylamino]propyl]phenoxy]-1-(4-carboxyphenoxy)-2-propanone

In the July 8, 2004 Office Action, the Examiner states that "Claim 11 will be allowed to the extent it reads on the elected subject matter. Compounds containing Silicon and heterocyclic subject matter should be deleted." Accordingly, Applicants have amended Claims 1, 2, 3, 7, 11, 12, and 13 to remove silicon and heterocyclic subject matter which reflects the scope of the generic concept of the elected subject matter. Applicants maintain the right to file divisional application(s) on non-elected subject matter.

Rejection of Claims 1-4, 9, 12, and 13 Under 35 U.S.C. §102(b)

The Examiner has rejected Claims 1-4, 9, 12, and 13 under 35 U.S.C. §102(b) as being unpatentable over JP 5-222006. JP'006 teaches compounds containing a

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heterocycle. It is Applicants' position that the amendments to Claims 1, 2, 3, 9, 12, and 13, which remove all heterocyclic subject matter, render the rejections moot. Therefore, it is respectfully requested that the rejections to Claims 1-4, 9, 12, and 13 be withdrawn.

Objection of Claim 7 and Allowance of Claim 11

Claim 7 has been objected to as being dependent upon a rejected base claim, but would be allowable to the extent that it reads on the elected subject matter, if rewritten in independent form including all of limitations of the base claim. The Examiner further states "Note applicants should delete Silicon containing subject matter." Applicants have amended Claim 7 to remove all silicon containing subject matter and respectfully request that the objection be withdrawn.

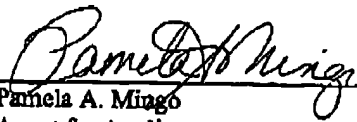
Claim 11 is allowed to the extent it reads on the elected subject matter. The Examiner further states "Compounds containing Silicon and heterocyclic subject matter should be deleted." Applicants have amended Claim 11 to remove all silicon and heterocyclic subject matter and respectfully request that the claim be allowed.

While Applicants submit that the claims are in condition for allowance and respectfully request the Examiner's reconsideration, a NOTICE OF APPEAL has nevertheless been filed. The Commissioner is hereby authorized to charge any additional fees under 37 CFR §1.17 which may be required, or credit any overpayment, to Account No. 19-3880 in the name of Bristol-Myers Squibb Company.

Respectfully submitted,

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Patent Department
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Date: October 7, 2004


Pamela A. Mingo
Agent for Applicants
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